AMENDMENTS TO THE CLAIMS

1. (Currently amended) A compound of Formula I or a stereoisomer thereof, or a pharmaceutically acceptable salt, or a prodrug thereof

$$R_1$$
 R_2 R_2 A_1

Formula I

wherein

X is selected from -NR₃R₄,-OR₃,-CR₃R₅R₅,-C(O)R₃,-S(O)_mR₃, NR₃C(O)R₄, or -NR₃S(O)_mR₄;

V is selected from -O-,-NR₅, or -S(O)_m;

m at each occurrence is 0, 1 or 2;

 R_1 and R_2 are independently selected from -NH(alkyl), -N(alkyl)₂, -NH(substituted alkyl), -N(substituted alkyl)₂, -O(alkyl), -O(substituted alkyl), halogen, alkyl, substituted alkyl, haloalkyl, cycloalkyl, substituted cycloalkyl, substituted phenyl, naphthyl, substituted naphthyl, heteroaryl, heteroaryl derivatives, substituted aryl, heterocycloalkyl, substituted heterocycloalkyl, substituted heteroaryl, -CR₅R₆Ar, -OAr, -S(O)_mAr, -NR₅Ar, -S(O)_malkyl, -S(O)_msubstituted alkyl, -NO₂, -OH, -NH₂, -SH, -C(O)NR₄R₅, -C(S)NR₄R₅, -C(O)NR₅Ar, -S(O)_mNR₅Ar, -S(O)_mNR₅Ar, -NR₅C(O)Ar, -NR₅S(O)_mAr, -NR₅S(O)_mAr, -C(O)Ar, -(alkyl linker)S(O)_mNH₂, -(alkyl linker)S(O)_mNR₅Ar, and -(alkyl linker)C(O)Ar;

R₃ and R₄ are independently selected from -H, alkyl, substituted alkyl, haloalkyl, cycloalkyl, substituted cycloalkyl, aryl, heterocycloalkyl, substituted heterocycloalkyl, substituted aryl cycloalkyl, substituted aryl cycloalkyl, heteroaryl cycloalkyl, substituted heteroaryl cycloalkyl, aryl heterocycloalkyl, substituted aryl heterocycloalkyl, heteroaryl heterocycloalkyl, or substituted heteroaryl heterocycloalkyl, the heterocycloalkyl, aryl heterocy

Each R₅ is independently selected from -H, alkyl, alkylene, alkylyne, cycloalkyl, haloalkyl, and alkyl substituted with 1-3 substituents selected from halogen, -O(alkyl), -NH(alkyl), -N(alkyl)₂, -C(O)NH(alkyl), -C(O)N(alkyl)₂, -

NHC(O)alkyl, -N(alkyl)C(O)alkyl, -S(O)_malkyl, heterocycloalkyl, substituted heterocycloalkyl and Ar. Ar;

Each R₆ is independently selected from alkyl, cycloalkyl, haloalkyl, and alkyl substituted with 1-3 substituents selected from halogen, -O(alkyl), -NH(alkyl), -N(alkyl), -C(O)N(alkyl)₂, -NHC(O)alkyl, -N(alkyl)C(O)alkyl, -S(O)_malkyl, heterocycloalkyl, substituted heterocycloalkyl and Ar;

provided that when V is –NH-; Ar is phenyl, substituted with methyl or nitro; R₁ is 2-(N,N-dimethylaminoethyl); R₂ is C(O)NH₂ then X is not CR₃R₅R₅.

Halogen is a group selected from-F,-Cl,-Br,-I;

Alkyl means both straight-and branched chain hydrocarbon chains having from 1-10 carbon atoms;

Alkylene means both straight-and branched chain hydrocarbon chains having from 2-10 carbon atoms and a double bond;

Alkylyne means both straight-and branched chain hydrocarbon chains having from 2-10 carbon atoms and a triple bond;

Substituted alkyl is an alkyl moiety from 1-10 carbon atoms having 1-3 substituents independently selected from halogen, $-S(O)_mR_5$, $-NR_5R_5$, $-C(O)R_5$, $-C(O)R_5$, $-C(O)R_5$, $-NR_5S(O)_mR_5$, $-NR_5S(O)_mR_5$, $-NR_5S(O)_mR_5$, and Ar;

Haloalkyl is an alkyl moiety having from 1-10 carbon atoms and having 1 to (2v+1) independently selected halogen substituent (s) where v is the number of carbon atoms in the moiety;

Cycloalkyl is a monocyclic or bicyclic alkyl moiety, having from 3-10 carbon atoms optionally containing 1 to 2 double bonds provided that the moiety is not aromatic, and further provided that the double bonds are not cumulated;

The term "substituted cycloalkyl" is a cycloalkyl group having 1-3 substituents independently selected from halogen,- R_5 , -OR $_5$, -S(O) $_mR_5$, -NR $_5R_5$, -C(O)R $_5$, -CN, -C(O)NR $_5R_5$, -NR $_5C$ (O)R $_5$, -S(O) $_mNR_5R_5$, -NR $_5S$ (O) $_mR_5$, and -NO $_2$;

Alkyl linker means a group selected from alkyl, substituted alkyl, haloalkyl, cycloalkyl, and substituted cycloalkyl having two points of attachment;

The term "heterocycloalkyl", unless otherwise specified, means a 4 to 8 membered monocylic ring or bicyclic ring, wherein at least one carbon atom is replaced with a heteromember selected from oxygen, nitrogen, -NH-, or -S(O)_m wherein m is zero, 1, or 2, optionally containing from one to three double bonds,

provided that the molecule is not aromatic; and provided that ring attachment can occur at either a carbon or nitrogen atom;

The term "substituted heterocycloalkyl" is a heterocycloalkyl group having 1-3 substituents independently selected from halogen, alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, $-OR_5$, $-S(O)_mR_5$, $-NR_5R_5$, $-C(O)R_5$, -CN, $-C(O)NR_5R_5$, $-NR_5C(O)R_5$, $-S(O)_mNR_5R_5$, $-NR_5S(O)_mR_5$, and $-NO_2$;

Substituted phenyl is a phenyl group having 1-3 substituents independently selected from halogen, alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, - OR_5 , - $S(O)_mR_5$, - NR_5R_5 , - $C(O)R_5$, - $C(O)R_5$, - $C(O)NR_5R_5$, - $NR_5C(O)R_5$, - $S(O)_mNR_5R_5$, - $NR_5C(O)R_5$, and - NO_2 ;

Substituted napthyl is a napthyl group having 1-3 substituents independently selected from halogen, alkyl, substituted alkyl, cycloalkyl, substituted cycloalkyl, - OR_5 , - $S(O)_mR_5$, - NR_5R_5 , - $C(O)R_5$, - $C(O)R_5$, - $C(O)NR_5R_5$, - $NR_5C(O)R_5$, - $S(O)_mNR_5R_5$, - $NR_5C(O)R_5$, and - NO_2 ;

The term "heteroaryl" means a radical attached via a ring carbon or nitrogen atom of a monocyclic aromatic ring containing five or six ring atoms consisting of carbon and 1, 2,3, or 4 heteroatoms each selected from the group consisting of non-peroxide O, S, N, with appropriate bonding to satisfy valence requirements as well as a radical (attachment at either carbon or nitrogen) of a fused bicyclic heteroaromatic of about eight to ten ring atoms;

The term "substituted heteroaryl" means a heteroaryl group having 1-3 substituents independently selected from halogen, $-R_5$, $-OR_5$, $-S(O)_mR_5$, $-NR_5R_5$, $-C(O)R_5$, $-C(O)NR_5R_5$, $-NR_5C(O)R_5$, $-S(O)_mNR_5R_5$, $-NR_5S(O)_mR_5$, and $-NO_2$, phenyl, substituted phenyl, napthyl, substituted napthyl, heteroaryl, and heteroaryl derivatives;

The term "heteroaryl derivatives" means a heteroaryl group having 1-3 substituents independently selected from halogen, -R₅, -OR₅, -S(O)_mR₅, -NR₅R₅, -C(O)R₅, -CN, -C(O)NR₅R₅, -NR₅C(O)R₅, -S(O)_mNR₅R₅, -NR₅S(O)_mR₅, and -NO₂;

Aryl is selected from phenyl, napthyl, substituted phenyl, substituted napthyl, heteroaryl, and substituted heteroaryl derivatives;

Ar is selected from aryl, substituted aryl, and substituted heteroaryl;

The term"aryl cycloalkyl"means a bicyclic ring system containing 9 to 14 carbon atoms wherein one ring is aryl and the other ring is fused to the aryl ring and may be fully or partially saturated in the portion of the ring not fused to the aryl ring, provided that either ring may act as a point of attachment;

The term "substituted aryl cycloalkyl" means an aryl cycloalkyl group having 1-3 substituents independently selected from halogen, - R_5 , - OR_6 , - $S(O)_mR_5$, - NR_5R_5 , - $C(O)R_5$, - $C(O)NR_5R_5$, - $NR_5C(O)R_5$, - $S(O)_mNR_5R_5$, - $NR_5S(O)_mR_5$, and - NO_2 ;

The term "heteroaryl cycloalkyl" means a bicyclic ring system containing 9 to 14 atoms, wherein one ring is heteroaryl and the other ring is fused to the aryl ring and may be fully or partially saturated in the portion of the ring not fused to the aryl ring, provided that either ring may act as a point of attachment;

The term "substituted heteroaryl cycloalkyl" means a heteroaryl cycloalkyl having 1-3 substituents independently selected from halogen, $-R_5$, $-OR_5$, $-S(O)_mR_5$, $-NR_5R_5$, $-C(O)R_5$, $-C(O)R_5$, $-C(O)NR_5R_5$, $-NR_5C(O)R_5$, $-S(O)_mNR_5R_5$, $-NR_5S(O)_mR_5$, and $-NO_2$;

The term "aryl heterocycloalkyl" means a bicyclic ring system containing 9 to 14 atoms, wherein one ring is aryl and the other ring is heterocycloalkyl, provided that either ring may act as a point of attachment;

The term "substituted aryl heterocycloalkyl" means an aryl heterocycloalkyl having 1-3 substituents independently selected from halogen, -R₅, -OR₅, -S(O)_mR₅, -NR₅R₅, -C(O)R₅, -C(O)NR₅, -NR₅C(O)R₅, -S(O)_mNR₅R₅, -NR₅S(O)_mR₅, and -NO₂:

The term "heteroaryl heterocycloalkyl" means a bicyclic ring system containing 9 to 14 atoms, wherein one ring is heteroaryl and the other ring is heterocycloalkyl, provided that either ring may act as a point of attachment;

The term "substituted heteroaryl heterocycloalkyl" means an heteroaryl heterocycloalkyl having 1-3 substituents independently selected from halogen, -R₅, -OR₅, -S(O)_mR₅, -NR₅R₅, -C(O)R₅, -CN, -C(O)NR₅R₅, -NR₅C(O)R₅, -S(O)_mNR₅R₅, -NR₅C(O)_mR₅, and -NO₂.

2. (Currently amended) A pharmaceutical composition comprising a compound according to Claim 1 or a pharmaceutically acceptable salt thereof and at least one pharmaceutically acceptable carrier or excipient.

3.-4. (Canceled)

- 5. (Currently amended) A <u>The</u> compound according to Claim 1 <u>or a</u> <u>pharmaceutically acceptable salt thereof</u> wherein the compound exhibits an IC50 for CRF binding of 1 micromolar or less.
- 6. (Currently amended) A <u>The</u> compound according to Claim 1 <u>or a</u> <u>pharmaceutically acceptable salt thereof</u> wherein the compound exhibits an IC50 for CRF binding of 100 nanomolar or less.
- 7. (Currently amended) A <u>The</u> compound according to Claim 1 <u>or a</u> <u>pharmaceutically acceptable salt thereof</u> wherein the compound exhibits an IC50 for CRF binding of 10 nanomolar or less in a standard assay of CRF binding.
- 8. (Currently amended) A method for treating stress, of treating anxiety or depression comprising administering to a patient in need thereof an a therapeutically effective amount of a compound according to Claim 1.
- 9. (Currently amended) A The compound according to claim 1 or a pharmaceutically acceptable salt thereof wherein V is O.
- 10. (Currently amended) A The compound according to claim 1 or a pharmaceutically acceptable salt thereof wherein V is NR₅.
- 11. (Currently amended) A The compound according to claim 1 or a pharmaceutically acceptable salt thereof wherein V is S.
- 12. (Currently amended) A The compound according to claim 1 or a pharmaceutically acceptable salt thereof wherein Ar is aryl.

- 13. (Currently amended) A The compound according to claim 1 or a pharmaceutically acceptable salt thereof wherein Ar is substituted aryl.
- 14. (Currently amended) A The compound according to claim 1 or a pharmaceutically acceptable salt thereof wherein Ar is substituted heteroaryl.
- 15. (Currently amended) A <u>The</u> compound according to claim 1 <u>or a</u> <u>pharmaceutically acceptable salt thereof</u> selected from the group consisting of (IR,2S)-1-({3,6-diethyl-5-[(4-methylpyridin-2-yl)oxy]pyrazin-2-yl}amino)-2,3-dihydro-1H-inden-2-ol,

N-[(1R,2S)-2-ethoxy-2,3-dihydro-1H-inden-1-yl]-3,6-diethyl-5-[(4-methylpyridin-2-yl)oxy]pyrazin-2-amine,

- 3,6-diethyl-N-[(1R,2S)-2-(2-fluoroethoxy)-2,3-dihydro-1H-inden-1-yl]-5-[(4-methylpyridin-2-yl)oxy]pyrazin-2-amine,
- 3,6-diethyl-N-[(IR,2S)-2-isopropoxy-2,3-dihydro-IH-inden-1-yl]-5-[(4-methylpyridin-2-yl)oxy]pyrazin-2-amine,
- 3,6-diethyl-5-[(4-methylpyridin-2-yl)oxy]-N-[(1R,2S)-2-propoxy-2,3-dihydro-IH-inden-1-yl]pyrazin-2-amine,
- (IR,2S)-1-({3,6-diethyl-5-[(4-methylpyridin-2-yl)oxy]pyrazin-2-yl}amino)-2,3-dihydro-IH-inden-2-yl acetate,
- (IR,2S)-1-({3,6-diethyl-5- [(4-ethylpyridin-2-yl)oxy] pyrazin-2-yl}amino)indan-2-ol,
- N-[(1R,2S)-2-ethoxy-2,3-dihydro-1H-inden-1-yl]-3,6-diethyl-5-[(4-ethylpyridin-2-yl)oxy]pyrazin-2-amine,
- (IR,2S)-I-({3,6-diethyl-5-[(3-methylpyridin-2-yl)oxyapyrazin-2-yl}amino)indan-2-ol,
- N-[(IR,2S)-2-ethoxy-2,3-dihydro-IH-inden-I-yl]-3,6-diethyl-5-[(3-methylpyridin-2-yl)oxy]pyrazin-2-amine,
- (1R,2S)-1-({3,6-diethyl-5-[(5-methylpyridin-2-yl)oxy]pyrazin-2-yl}amino)indan-2-ol, N-[(1R,2S)-2-ethoxy-2,3-dihydro-IH-inden-l-yl]-3,6-diethyl-5-[(5-methylpyridin-2-yl)oxy]pyrazin-2-amine,

5-[(4,6-dimethylpyridin-2-yl)oxy]-N-[(1R,2S)-2-ethoxy-2,3-dihydro-IH-inden-1-yl]-3,6-diethylpyrazin-2-amine,

N-[(IR,2S)-2-ethOxy-2,3-dihydro-IH-inden-I-yl]-3,6-diethyl-5-(3-methylphenoxy)-pyrazin-2-amine,

1-({3,6-diethyl-5-[(4-methylphenyl)amino]pyrazin-2-yl}amino)indan-2-ol,

N-(2-ethoxy-2,3-dihydro-lH-inden-l-yl)-3,6-diethyl-5-[(4-methylphenyl)thio] pyrazin-2-amine,

3,6-diethyl-N-[(IR,2S)-2-(2-fluoroethoxy)-2,3-dihydro-IH-inden-l-yl]-5-[(4methylpyridin-2-yl)oxy]pyrazin-2-amine, and

N-[(1R,2S)-2-ethoxy-2,3-dihydro-1H-inden-1-yl]-3,6-diethyl-5-[(4-methylpyridin-2-yl)oxy]pyrazin-2-amine.